

AMENDMENTS TO THE SPECIFICATION

IN THE SPECIFICATION:

Page 6

The paragraph beginning on page 6, line 28, and ending on page 7, line 5, is being amended as follows:

the C1-C6 haloalkoxy group includes, for example, a C1-C4 haloalkoxy group such as a fluoromethoxy group, a difluoromethoxy group, a trifluoromethoxy group, ~~a chloromethyl group, a dichloromethyl group, a trichloromethyl group,~~ a 2,2,2-trifluoroethoxy group, a 1,1,2,2,-tetrafluoroethoxy group, a 2-fluoroethoxy group and the like;

Page 7

The paragraph beginning on page 7, line 16, is being amended as follows:

the (C1-C6 alkylsulfonyloxy) C1-C6 alkyl group includes a methylsulfonyloxymethyl group, a ethylsulfonyloxymethyl group, a 1-methylsulfonyloxyethyl group, ~~a 2-sulfonyloxyethyl group,~~ a 1-methylsulfonyloxypropyl group;

Page 14

The paragraph beginning on page 14, line 11, is being amended as follows:

the C1-C4 haloalkoxy group includes, for example, a fluoromethoxy group, a difluoromethoxy group, a trifluoromethoxy group, a chloromethyl group, a dichloromethyl group, a trichloromethyl group, a 2,2,2-trifluoroethoxy group, a 1,1,2,2-tetrafluoroethoxy group, a 2-fluoroethoxy group;

the di(C1-C4alkyl)amino group includes, for example, a dimethylamino group, a diethylamino group, a dipropylamino group;

Page 68

At page 68, please delete the paragraph starting at line 2 and extending to line 7, in its entirety.

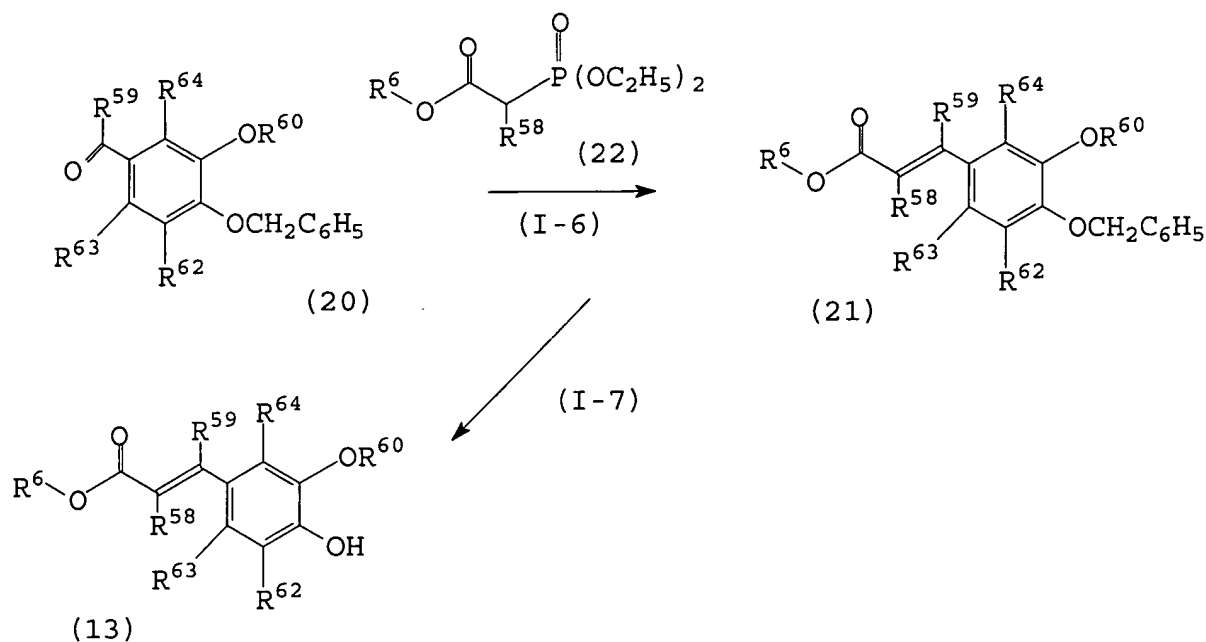
Page 72

The paragraph beginning on page 72, line 9, is being amended as follows:

The compound ~~(11)~~ (13) can be produced by subjecting the compound (12) to hydrolysis reaction in the presence of an acid.

Page 77

Please amend the Specification beginning on page 77, line 1, as follows:



wherein, in the formula,

R^6 represents a C1-C6 alkyl group, ~~R^7 represents a benzyl group or a methoxymethyl group,~~ R^{58} , R^{59} , R^{60} , R^{62} , R^{63} , R^{64} , L^1 and L^3 represent same meaning as described above.

Page 78

The paragraph beginning on page 78, line 9, is being amended as follows:

The compound (13) can be produced by making the compound ~~(20)~~ (21) react with hydrogen in the presence of a hydrogenation catalyst and an acid.

Page 103

The paragraph beginning on page 103, line 4, is being amended as follows:

4-fluorophenyl group, 4-chlorophenyl group, 4-bromophenyl group, 4-iodophenyl group, 4-methylphenyl group, 4-ethylphenyl group, 4-propylphenyl group, 4-isopropylphenyl group, 4-butylphenyl group, 4-(sec-butyl)phenyl group, 4-isobutylphenyl group, 4-(tert-butyl)phenyl group, 4-vinylphenyl group, 4-(1-methylvinyl)phenyl group, 4-(1-propenyl)phenyl group, 4-(2-methyl-1-propenyl)phenyl group, 4-(1-butenyl)phenyl group, 4-ethynylphenyl group, 4-(1-propynyl)phenyl group, 4-(1-butynyl)phenyl group, ~~4-(1-pentanyl)phenyl group~~, 4-(3-methyl-1-butynyl)phenyl group, 4-(3,3-dimethyl-1-butynyl)phenyl group, 4-(fluoromethyl)phenyl group, 4-(difluoromethyl)phenyl group, 4-(trifluoromethyl)phenyl group, 4-methoxyphenyl group, 4-ethoxyphenyl group, 4-(fluoromethoxy)phenyl group, 4-(difluoromethoxy)phenyl group, 4-(trifluoromethoxy)phenyl group, 4-cyanophenyl group, 4-(N,N-dimethylamino)phenyl group, 4-(N,N-diethylamino)phenyl group, 4-(N,N-dipropylamino)phenyl group, 3,4-difluorophenyl group, 4-chloro-3-fluorophenyl group, 4-bromo-3-fluorophenyl group, 3-fluoro-4-methylphenyl group, 4-ethyl-3-fluorophenyl group, 3-fluoro-4-(trifluoromethyl)phenyl group, 3-fluoro-4-methoxyphenyl group, 4-cyano-3-fluorophenyl

group, 4-fluoro-3-chlorophenyl group, 3,4-dichlorophenyl group, 4-bromo-3-chlorophenyl group, 3-chloro-4-methylphenyl group, 3-chloro-4-ethylphenyl group, 3-chloro-4-(trifluoromethyl)phenyl group, 3-chloro-4-methoxyphenyl group, 4-cyano-3-chlorophenyl group, 3-bromo-4-fluorophenyl group, 3-bromo-4-chlorophenyl group, 3,4-dibromophenyl group, 3-bromo-4-methylphenyl group, 3-bromo-4-ethylphenyl group, 3-bromo-4-(trifluoromethyl)phenyl group, 3-bromo-4-methoxyphenyl group, 3-bromo-4-cyanophenyl group, 4-fluoro-3-methylphenyl group, 4-chloro-3-methylphenyl group, 4-chloro-3-methylphenyl group, 3,4-dimethylphenyl group, 4-ethyl-3-methylphenyl group, 3-methyl-4-(trifluoromethyl)phenyl group, 4-methoxy-3-methylphenyl group, 4-cyano-3-methylphenyl group, ~~3-ethyl-4-fluoromethyl group~~, 4-chloro-3-ethylphenyl group, 4-bromo-3-ethylphenyl group, 3-ethyl-4-methylphenyl group, 3,4-diethylphenyl group, 3-ethyl-4-(trifluoromethyl)phenyl group, 3-ethyl-4-methoxyphenyl group, 4-cyano-3-ethylphenyl group, 4-fluoro-3-(trifluoromethyl)phenyl group, 4-chloro-3-(trifluoromethyl)phenyl group, 4-bromo-3-(trifluoromethyl)phenyl group, 4-methyl-3-(trifluoromethyl)phenyl group, 4-ethyl-3-(trifluoromethyl)phenyl group, 3,4-di-(trifluoromethyl)phenyl group, 4-methoxy-3-(trifluoromethyl)phenyl group, 4-cyano-3-(trifluoromethyl)phenyl group, 4-methoxycarbonylphenyl group, 4-ethoxycarbonylphenyl

group, 4-phenylphenyl group, 3-fluoro-4-phenylphenyl group, 3-chloro-4-phenylphenyl group, 3-bromo-4-phenylphenyl group, 3-methyl-4-phenylphenyl group, 3-ethyl-4-phenylphenyl group, 4-phenoxyphenyl group, 3-fluoro-4-phenoxyphenyl group, 3-chloro-4-phenoxyphenyl group, 3-bromo-4-phenoxyphenyl group, 3-methyl-4-phenoxyphenyl group, 3-ethyl-4-phenoxyphenyl group, 4-nitrophenyl group, 3-fluoro-4-nitrophenyl group, 3-chloro-4-nitrophenyl group, 3-bromo-4-nitrophenyl group, 3-methyl-4-nitrophenyl group, 3-ethyl-4-nitrophenyl group, 4-methylthiophenyl group, 3-fluoro-4-methylthiophenyl group, 3-chloro-4-methylthiophenyl group, 3-bromo-4-methylthiophenyl group, 3-methyl-4-methylthiophenyl group, 3-ethyl-4-methylthiophenyl group,

Indan-5-yl group, 5,6,7,8-tetrahydronaphthalene-2-yl group, 6,7,8,9-tetrahydro-5H-benzocycloheptene-2-yl group, 5,6,7,8,9,10-hexahydro-benzocyclooctene-2-yl group, 2-naphthyl group, 4-fluoronaphthalene-2-yl group, 5-fluoronaphthalene-2-yl group, 6-fluoronaphthalene-2-yl group, 7-fluoronaphthalene-2-yl group, 4-chloronaphthalene-2-yl group, 5-chloronaphthalene-2-yl group, 6-chloronaphthalene-2-yl group, 7-chloronaphthalene-2-yl group, 4-bromonaphthalene-2-yl group, 5-bromonaphthalene-2-yl group, 6-bromonaphthalene-2-yl group, 7-bromonaphthalene-2-yl group, 4-methylnaphthalene-2-yl group, 5-methylnaphthalene-2-yl

group, 6-methylnaphthalene-2-yl group, 7-methylnaphthalene-2-yl group, 4-methoxynaphthalene-2-yl group, 5-methoxynaphthalene-2-yl group, 6-methoxynaphthalene-2-yl group, 7-methoxynaphthalene-2-yl group, 4-trifluoromethylnaphthalene-2-yl group, 5-trifluoromethylnaphthalene-2-yl group, 6-trifluoromethylnaphthalene-2-yl group, 7-trifluoromethylnaphthalene-2-yl group, 5,6-difluoronaphthalene-2-yl group, 5,6-dichloronaphthalene-2-yl group, 5,6-dimethylnaphthalene-2-yl group, 5-fluoro-6-methylnaphthalene-2-yl group, 6-fluoro-5-methylnaphthalene-2-yl group, 5-chloro-6-methylnaphthalene-2-yl group, 6-chloro-5-methylnaphthalene-2-yl group, 6-chloro-5-fluoronaphthalene-2-yl group, 5-chloro-6-fluoronaphthalene-2-yl group, 4-(2-fluorovinyl)phenyl group, 4-(2-chlorovinyl)phenyl group, 4-(2-bromovinyl)phenyl group, 4-(2,2-difluorovinyl)phenyl group, 4-(2,2-dichlorovinyl)phenyl group, 4-(2,2-dibromovinyl)phenyl group, 4-(1-methyl-2,2-dichlorovinyl)phenyl group, 4-(1-methyl-2,2-dibromovinyl)phenyl group, 4-chloroethynylphenyl group, 4-bromoethynylphenyl group, 4-iodoethynylphenyl group, 4-methoxymethylphenyl group, 4-ethoxymethylphenyl group, 4-propyloxymethylphenyl group, 4-(1-methoxyethyl)phenyl group, 4-(2-methoxyethyl)phenyl group, 4-(2-methoxypropyl)phenyl group, 4-(2-methoxyisopropyl)phenyl group, 4-phenoxyethylphenyl group,

4-(1-phenoxyethyl)phenyl group, 4-(2-phenoxyethyl)phenyl group,
4-(1-phenoxypropyl)phenyl group, 4-(3-phenoxypropyl)phenyl
group, 4-(4-phenoxybutyl)phenyl group, 4-(hydroxymethyl)phenyl
group, 4-(1-hydroxyethyl)phenyl group, 4-(2-hydroxyethyl)phenyl
group, 4-(1-hydroxypropyl)phenyl group, 4-(1-hydroxybutyl)phenyl
group, ~~4-methylsulfonyloxymethyl)phenyl group, 4-~~
~~ethylsulfonyloxymethyl)phenyl group~~ 4-
(methylsulfonyloxymethyl)phenyl group, 4-
(ethylsulfonyloxymethyl)phenyl group, 4-(1-
~~methylsulfonyloxyethyl)phenyl group, 4-(2-~~
~~sulfonyloxyethyl)phenyl group, 4-(1-~~
~~methylsulfonyloxypropyl)phenyl group, 4-carbonylphenyl group, 4-~~
methylcarbonylphenyl group, 4-ethylcarbonylphenyl group, 4-
propylcarbonylphenyl group, 4-isopropylcarbonylphenyl group, 4-
methoxyiminomethylphenyl group, 4-(1-methoxyiminoethyl)phenyl
group, 4-(1-methoxyiminopropyl)phenyl group, 4-(1-
methoxyiminobutyl)phenyl group, 4-ethoxyiminomethylphenyl group,
4-(1-ethoxyiminoethyl)phenyl group, 4-propoxyiminomethylphenyl
group, 4-(1-isopropoxyiminoethyl)phenyl group, 4-
(butoxyiminomethyl)phenyl group, 4-pentyloxyiminomethylphenyl
group, 4-(1-hexyloxyiminoethyl)phenyl group, 4-
benzyloxyiminomethylphenyl group, 4-(1-
benzyloxyiminoethyl)phenyl group, 4-(1-

benzyloxyiminopropyl)phenyl group, 4-(1-benzyloxyiminobutyl)phenyl group, 4-dimethylaminoiminomethylphenyl group, 4-(1-dimethylaminoiminoethyl)phenyl group, 4-diethylaminoiminomethylphenyl group, 4-(1-diethylaminoiminoethyl)phenyl group, 4-trimethylsilylphenyl group, 4-triethylsilylphenyl group, 4-tert-butyl dimethylsilylphenyl group.

Page 120

The paragraph beginning on page 120, line 20, is being amended as follows:

By using 0.5 g of 3-{3-methoxy-4-(2-propynyloxy)phenyl}propionyl chloride, 0.42 g of 4-chloro-3-(trifluoromethyl)benzylamine and 0.42 ml of triethylamine according to the Production Example 18 was obtained 0.76 g of ~~N-{4-chloro-3-(trifluoromethyl)benzyl}-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide~~ N-{4-chloro-3-(trifluoromethyl)benzyl}-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide (referred as the compound of the present invention 19 hereinafter).

Page 147

The paragraph beginning on page 147, line 24, is being amended as follows:

In the same way as in the Production Example 40, 345 mg of ~~N-(3,4-dimethylbenzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanthioamide}~~ N-(3,4-dimethylbenzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanthioamide (referred as the compound of the present invention 59 hereinafter) was obtained from 0.43 g of N-(3,4-dimethylbenzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide and 0.36 g of Lawesson's Reagent.

Page 162

The paragraph beginning on page 162, line 14, is being amended as follows:

Production Example 82

538 mg of N-(4-chlorobenzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide and 5 ml of N,N-dimethylformamide were mixed and cooled to 0 °C. 72 mg of sodium hydride was added to the mixture, and it was stirred at 0 °C for about 15 minutes. Then, 196 mg of propargyl bromide was added in it, the mixture was stirred at 0 °C for 30 minutes and at room temperature for 2 hours. After that, water was added to the reaction mixture and extracted with ethyl acetate. The organic

layer was washed successively with 5 % hydrochloric acid, saturated aqueous solution of sodium bicarbonate and saturated aqueous solution of sodium chloride, dried by magnesium sulfate, and concentrated under reduced pressure. The residue was purified by silica gel column to obtain 105 mg of ~~N-(4-chlorobenzyl)-N-propynyl-3-(4-methoxy-3-(2-propynyloxy)phenyl}propanamide~~ N-(4-chlorobenzyl)-N-propynyl-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide (referred as the compound of the present invention 82 hereinafter).

Page 172

The paragraph beginning on page 172, line 12, is being amended as follows:

By using 0.30 g of 3-{3-methoxy-4-(2-propynyloxy)phenyl}propionic acid and 0.22 g of 1-(5,6,7,8-tetrahydronaphthalene-2-yl)ethylamine according to the Production Example 94 was obtained 0.22 g of N-{1-(5,6,7,8-tetrahydronaphthalene-2-yl)ethyl}3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide (referred as the compound of the present invention ~~97~~ 96 hereinafter).

Page 173

The paragraph beginning on page 173, line 3, is being amended as follows:

By using 0.30 g of 3-{3-fluoro-5-methoxy-4-(2-propynyloxy)phenyl}propionic acid and 0.22 g of 4-chlorobenzylamine according to the Production Example 94 was obtained 0.23 g of ~~N-(4-chlorobenzyl)-3-{3-fluoro-5-methoxy-(2-propynyloxy)phenyl}propanamide~~ N-(4-chlorobenzyl)-3-{3-fluoro-5-methoxy-4-(2-propynyloxy)phenyl}propanamide (referred as the compound of the present invention 97 hereinafter).

Page 178

The paragraph beginning on page 178, line 6, is being amended as follows:

0.50 g of N-(4-methylcarbonylbenzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide, 0.21 ml of 1,1-dimethylhydrazine, 1ml of acetic acid and 10 ml of methanol were mixed, and stirred under the condition of reflux for 1 hour. Then water was added to the reaction mixture, and extracted with ethyl acetate. The organic layer was washed successively with 5 % hydrochloric acid twice, saturated aqueous solution of sodium bicarbonate and saturated aqueous solution of sodium chloride, dried by magnesium sulfate, and concentrated under reduced pressure. The obtained solid was washed by hexane to obtain

0.46 g of ~~N-(4-(1-dimethylaminoimino)ethyl)benzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide~~ N-(4-(1-dimethylaminoiminoethyl)benzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide (referred as the compound of the present invention 104 hereinafter).

Page 220

The paragraph beginning on page 220, line 16, is being amended as follows:

1.59 g of 4-hydroxy-benzylamine hydrochloride, 2.34 g of 3-{3-methoxy-4-(2-propynyloxy)phenyl}-propionic acid, 2.11 g of WSC and 25 ml of pyridine were mixed, and stirred at room temperature for 4 hours. Then water was added to the reaction mixture and extracted with ethyl acetate. The organic layer was washed successively with 5 % hydrochloric acid, water and saturated aqueous solution of sodium chloride, dried by magnesium sulfate, and concentrated under reduced pressure. The residue was purified by silica gel column to obtain 2.1 g of ~~N-(4-hydroxy-benzyl)-3-{4-methoxy-3-(2-propynyloxy)phenyl}propanamide~~ N-(4-hydroxy-benzyl)-3-{3-methoxy-4-(2-propynyloxy)phenyl}propanamide.

Page 248

The paragraph beginning on page 248, line 7, is being amended as follows:

According to Synlett, 2000, No.12, p1801 to 1803, 8.0 g of 4-bromobenzonitrile, trisdibenzylideneacetonepalladium(0), 0.98 g of 2-(di-tert-butylphosphino)biphenyl and 200 ml of dimethylimidazoline were mixed at room temperature for 5 minutes hours. Then 16.9 ml of hexamethyldisilane and 2.0 g of water were added to it and washed with water for five times, dried by magnesium sulfate, concentrated under reduced pressure. The residue was purified by silica gel column to obtain 8.0 g of 4-trimethylsilylbenzonitrile was obtained from 10 g of 4-bromobenzonitrile, trisdibenzylideneacetonepalladium(0), 0.98 g of 2-(di-tert-butylphosphino)biphenyl and 200 ml of dimethylimidazoline.